Amendments to the Specification:

Please replace paragraph [0668] of the published application, with the following amended paragraph:

Error-prone frequency was regulated by changing the proofreading function of DNA polymerase δ or ε. The proofreading function was changed by producing disparity mutant strains which had a deletion in the proofreading portion of DNA polymerase δ or ε. To produce mutant strains, site-directed mutagenesis was used to perform base substitutions at a specific site of DNA polymerases polδ or polε of the normal strain (Morrison A. & Sugino A., Mol. Gen. Genet. (1994) 242: 289-296) using common techniques (Sambrook et al., Molecular Cloning: A Laboratory Manual, Ver. 2, Cold Spring Harbor Laboratory (Cold Spring Harbor, N.Y., 1989), supra). Specifically, conversion was performed in polδ ~~322~~ 321(D)→(A); and ~~324~~ 323(E)→(A); and in polε ~~291~~ 290(D)→(A); and ~~293~~ 292(E)→(A). These mutants were a DNA polymerase δ mutant strain (AMY128-1: Pol3-01 MATα, ura3-52leu2-1 lys1-1 ade2-1 his1-7 hom3-10 trp-289 canR; available from Prof. Sugino (Osaka University) and a DNA polymerase ε mutant strain (AMY2-6: pol2-4 MATα, ura3-52 leu2-1 lys1-1 ade2-6 his1-7 hom3-10 try1-289 canR; available from Prof. Sugino (Osaka University). It will be understood that equivalents of such strains can be produced by those skilled in the art using site directed mutagenesis to introduce mutations, such as ~~322~~ 321(D)→(A) and ~~324~~ 323(E)→(A) in polδ; and ~~291~~ 290(D)→(A) and ~~293~~ 292(E)→(A) in polε.

Please replace paragraph [0727] of the published application with the following paragraph:

For Pol mutations, one-base mutation was introduced into the proofreading activity sites (SEQ ID Nos. 55 and 56 (δ); SEQ ID Nos. 57 and 58(ε)) of both polδ and polε to delete proofreading activity: in polδ ~~315~~ 314(D)→(A), ~~317~~ 316(E)→(A); and in polε 275(D)→(A), 277(E)→(A) (Morrison A. & Sugino A., Mol. Gen. Genet. 242: 289-296, 1994; Goldsby R.E., et al., Proc. Natl. Acad. Sci. USA, 99: 15560-15565, 2002).

Please replace paragraph [0745] of the published application, with the following paragraph:

Next similar experiments were carried out using rats as models. Rat models of cancer can be rapidly prepared by introducing mutations into polδ (in an amino acid sequence as set forth in SEQ ID NO. 60, D at position ~~315~~ 312 and E at position ~~317~~ 314 are substituted with alanine).